Attorney Docket No. 1324

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant: Mark David Hoffbeck

Date:

≃Dec. 10, 2002

Serial No.: 09/758,858 =

Group Art Unit:

638

Filed:

January 11, 2001 _- -

Examiner:

⇒David T. Fox

For.≪

"INBRED MAIZE LINE PH6WR"

Assistant Commissioner for Patents Washington, D.C. 20231

RÜLE 132 DECLARATION COF DR. STEPHEN SMITH

Sir:

- I, Stephen Smith, PhD., do hereby declare and say as follows:
- 1. I am skilled in the art of the field of the invention. I have a Ph.D. in Biochemical Systematics and Taxonomy of Maize and its Wild Relatives from Birmingham University. I have a M.Sc. in the Conservation and Utilization of Plant Genetic Resources from Birmingham University. I have a Bachelor of Science degree in Plant Sciences from London University. Since 1977 I have been engaged in the development, study and application of molecular markers to genetics, measuring genetic diversity and tracking pedigrees. I commenced this work at North Carolina State University as a post-doctoral research fellow. I have continued my engagement in these studies during my employment by Pioneer Hi-Bred from 1980 until the present. These studies have resulted in numerous scientific articles that have appeared in peer reviewed scientific literature.
 - 2. I have read and understood the Office Action in the above case dated October 30, 2002. This declaration is in response to the Examiner's rejection under, 35 U.S.C. § 102(e) as anticipated by or, in the afternative, under 35 U.S.C. § 103(a) as obvious over Morgan (U.S. Patent No. 5,824,848).
 - 3. I have conducted an analysis of SSR marker data for inbred PH6WR and-the inbred cited as prior art, F361. Out of a total of 68 SSR loci examined, which allowed a sampling of each chromosome, there are 41 markers that show differences between PH6WR and F361. This represents a difference for 60% for the markers tested. Of

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these 41 markers, 21 were greater than 50 cM in distance, or unlinked on the genetic map.

- 4. Upon crossing PH6WR to any other maize line and selfing successive filial generations, one would within the realm of what is statistically possible, obtain a progeny inbred maize-line that retains genetic contribution from PH6WR. Assuming that (i) the cited prior art is used as the maize line to which PH6WR is crossed, (ii) that the only difference between PH6WR and F361 are these 41 markers, and (iii) that all markers within a 50 cM distance will segregate together, then the odds of obtaining a PH6WR progeny inbred that is the same as F361 after one cycle of breeding, is 1 in 2²¹ or 1 in 2,097,152. Statistically it is extremely unlikely that a PH6WR progeny, after one cycle of breeding, would be the same as F361.
- PH6WR from F361. For example, it is common practice in quantitative genetics to determine the relation of plants by differences in markers. In doing so, one extrapolates that a percentage difference in markers is indicative of a difference in the whole genome. To assume that the only differences between PH6WR and F361 are for these 41 markers, when 41 markers constitute 60% of the 68 SSR loci examined, is a gross and unrealistic assumption. Further the current maize genetic map only has approximately sixty 50cM units, so by applying this limitation the maximum number of independently segregating loci one could obtain, using the most different maize lines that could ever be found, is sixty. These assumptions result in an over estimate of the odds of breeding PH6WR from F361.
 - 6. Given the difference in molecular markers between PH6WR and F361, it is my expert opinion that PH6WR and F361 are very distinct inventions. It is also my expert opinion that, within the realm of what is statistically possible, any progeny of PH6WR developed through crossing PH6WR with another plant will be distinct from F361. Given the facts and based on my education and scientific experience. I believe that the invention as claimed is not obvious nor anticipated by Morgan (U.S. Patent No. 5,824,848):
 - 7. I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

Date: <u>Dec 11 2002</u>

By: Neysh Mon, the

Stephen Smith

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Attorney Docket No. 1324

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant:

Mark David Hoffbeck

Date:

February 28, 2003 00

Serial No.:_

.09/758,858

-Group Art Unit:

1638

Filed:

January 11, 2001

Examiner:

David T. Fox

For:

"INBRED MAIZE-LINE PH6WR"

Assistant Commissioner for Patents Washington, D.C. 20231

RULE 132 DECLARATION
OFDR. DINAKAR BHATTRAMAKKI

Sir.

- I, Dinakar Bhattramakki, Ph.D., do hereby declare and say as follows:
- 1. I am skilled in the art of the field of the invention. I have a Ph.D. in Plant Molecular Genetics from the University of Illinois at Urbana-Champaign. I have a Bachelor of Science degree in Agricultural Sciences from the University of Agricultural Sciences, Bangalore, India. Since 1997 I have been engaged in the analysis of molecular markers for plants. I have supervised the Molecular Marker Applications lab at Pioneer Hi-Bred International, Inc. from January 2002 until the present.
- 2. I am familiar with the methods used in the analysis of Simple Sequence-Repeat, SSR, marker data for inbred PH6WR conducted at Rioneer Hi-Bred International, Inc. The analysis of the SSR profile of inbred PH6WR may be accomplished without any undue experimentation. The SSR profile for inbred PH6WR-is attached hereto.
- Means of performing this genetic marker profile are well known in the art. SSRs are genetic markers based on polymorphisms in nucleotide sequences. The PCR™ detection of SSRs is accomplished by using two oligonucleotide primers flanking the polymorphic segment of DNA. Amplification is accomplished through repeated cycles of heat denaturation of the DNA followed by annealing of the primers to their complementary sequences at low temperatures, and extension of the annealed primers with DNA polymerase.
- 4. Markers are scored following amplification and gel electrophoresis of the amplification products. Scoring of marker genotype is based on the size or weight of the amplified

Appendix D

fragment. While variation in the primer used or in laboratory procedures can affect the reported marker score, relative values remain constant regardless of the specific primer or laboratory used.

- 5. Primers that may be used to identify the SSR markers reported herein are publicly available and may be found in the Maize DB on the World Wide Web at agron.missouri.edu/maps.html (sponsored by the University of Missouri), in Sharopova et al. (Plant Mol. Biol. 48(5-6):463-481) and/or in Lee et al (Plant Mol. Biol. 48(5-6); 453-461). Markers shown for PH6WR are the publicly available markers in the sources listed above for which PH6WR was tested and shown to be homozygous.
- 6. Map information is provided by bin number as reported in the Maize DB. The bin number digits to the left of decimal point typically represent the chromosome on which such marker is located, and the digits to the right of the decimal typically represent the location on such chromosome.
- 7. I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

Date: February 28, 2003

Dinakar Bhattramakki

• .	2.7	:
Public Name	PH6WR	
😽 of Marker	Bin # base pairs	
phi427913	_{್ಸ್} 1.01 _උ ე129;38 ~	
bnlg1014	1.01 124.16	<
phi056	1.01 - 255.30	
bnlg1083	1.02 221.86	_
- bnig1127	1.023 96.21	
bnlg1429	1.02 - 190.09	_
bnlg1627_	1.02 198.87	
bnlg439	1.03 232.00	
phi109275	-1.03 131.94	-
phi339017	1.03 145.56	
bnlg1203	1.03 306.51	
bnlg1484-	1.03. 143.49 1.05 141.48	
bnlg1886 **	-4.C.	
bnlg1057 _bnlg1041	1.06 274.00 1.06 194.58	
bnig1615	1.06 - 211.96	
bnig 1515 bnig 1556	1.07 203.76	
phi323065	1:08 329.53	
- phi335539	1.08 88.58	
phi423298	1.08 133.68	
phi002	1.08 69.78	
bnlg1331	4.09 121.03	-
phi011	1.09 226.98	
phi227562	1.11 319.44	
phi265454	1.11 217.66	
phi064	1.11 - 84.29	
phi402893	2.00 212.91	
_bnig1017	2.02 _195.57	
bnlg2277	- 2.02 - 294.16	
bnlg1064	2.03 188.18	
bnlg1018	2.04. <u>1</u> 38.37	
bnlg1138	2.06 223.89	
bnlg1831	2.06 - 189.90	
phi251315	2.07123.79	
phi435417	2.08 214.27	
™ bînlg1141	2.08 152.92 2.08 122.04 -	
-phi127	-	
Tphi101049	•	
phi453121 phi104127	3.00 217.80 _ 3.01 169.96	. 12
phi404206	3:01 300.24	
phi193225	3.02 133.45	
phi374118	3.02 133.43	
bnlg1144	3.02 157.89	
bnlg1647	3.02 131.00	
ag.a,		

bnlg1523	3.03	. 318.23
bnlg1113	3.04	77.84
bnlg1452	3.04	96.77
bnlg1035 🔙	3.05	112.85
phi053	3.05	191.64
phi102228	3.06 -	- 135.04
bnlg1160 - 🤝	3.06	222.23
bnlg1951	- 3.06	126.99
bnlg2241 -	3.06	142.63
phi072	4.00	439.43
phi295450	4.01	184.83
bnlg1162	4.03	126.15
phi308090	4.04	216.06
=phi096.	4.04	234.58
phi438301	4.05	208.45
bnlg1159 -	~ 4.05	147.81 **
bnlg1755	4.05	216.63
bnlg1937 -	4.05	227.36
bnlg1265	4.05	198.45
phi079	4.05	177.74
bnlg1189	4.07	129.65
*bnlg2244	4.08	197.80
bnlg1006	5.00	227.74
phi396160	5.02	298,12
phi109188	5.03	161.52
bnig653	5.04	151.67
phi330507	5.04	131.59
~phi331888	5.04	134.18
bnlg1208	5.04,	122.87
~ bnlg1892	5.04	158.09
phi333597	5.05	210.74~
bnlg1118	5.07	72.54
phi423796	~ 6.01	128.39
~phi389203	6.03	306.15
phi452693	6.04	-123.30
_phi445613	- 6.05	100.12
bnlg1174 =	~6.05	218.72
phi299852	-6.07	119.55
phi364545-	6.07 ^{***}	131.46
bnlg1740	6.07	236.30
bnlg1759	6:07	136.00
bnlg2271-	7.03	219.98
phi328175	7.04	121.92
phi260485	7.05	285.39
phi051	7.05	136.61
phì116	7.06	168.82
phi420701	8.00	291.88
bnlg1194	8.02	140.77
• •		

phi100175	8.03	144.28
bnlg2082	8.031	172.52
phi121=	8.03	97.90
bnlg2046-	8.04	320.35
bnlg1176	8.05	194.15
້ bnig1152 🧠	8.06	121.65
_ bnlg1065	~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~	215.69
bnlg1056 -	8.08	108.46
phi015 -	8.08_	81789
phi233376	8.09	_ 147.93
-bnlg2122	9.01	234.97-
bnlg1012	₌ 9.04	~ 157.54
° phi032	9:04	236.63
phi108411	9.05 -	126.04
phi236654	9.05 ==	117.08
- bnlg1375 - "	9.07	164.93
_ bnlg1129 _	9.08	300.56
phi96342	10.02	252.35
- phi059	10.02	153.35
⁻ bnlg1655	10.03	148.23
phi050	10:03	83.27
phiD62	10.04	161.07.
phi323152	10.05	134.72
bnlg1074	10.05	161.94
bnlg1185	10:07	151:61
bnlg1450 .	10.07	189.35
phi448880	9.06/9.07	186.11